

FORM PTO-1449/A and B (modified PTO/SB/08) INFORMATION DISCLOSURE STATEMENT BY APPLICANT			APPLICATION NO.:	10/821,813	ATTY. DOCKET NO.:	P0453.70112US01
			FILING DATE:	April 8, 2004	CONFIRMATION NO.:	9059
			APPLICANT:	Boyd et al.		
			GROUP ART UNIT:	1614	EXAMINER:	SPIVACK

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U.S. PATENT DOCUMENTS

Examiner's Initials	Cite No.	U.S. Patent Document		Name of Patentee or Applicant of Cited Document	Date of Publication or Issue of Cited Document MM-DD-YYYY
		Number	Kind Code		
PS		4,322,426		Hermann et al.	03-30-1982
		4,965,269		Brändström et al.	10-23-1990
		5,202,159		Chen et al.	04-13-1993
		5,391,372		Campbell	02-21-1995
		5,567,423		Ying et al.	10-22-1996
		5,614,219		Wunderlich et al.	03-25-1997
		5,656,290		Kelm et al.	08-12-1997
		5,981,185		Matson et al.	11-09-1999
		6,455,537		Cooper	09-24-2002
		2002-0064771	A1	Zhong et al.	05-30-2002
		2003-0026801	A1	Weiner et al.	02-06-2003
		2003-0065003	A1	Foss et al.	04-03-2003
		2003-0187010	A1	Foss et al.	10-02-2003
		2004-0162306	A1	Foss et al.	08-19-2004
		2004-0162307	A1	Foss et al.	08-19-2004
		2004-0162308	A1	Foss et al.	08-19-2004
		2004-0167147	A1	Foss et al.	08-26-2004
		2004-0167148	A1	Foss et al.	08-26-2004
		2004-0259899	A1	Sanghvi et al.	12-23-2004
		2004-0266806	A1	Sanghvi et al.	12-30-2004
		2005-0004155	A1	Boyd et al.	01-06-2005
		2005-0048117	A1	Foss et al.	03-03-2005
		2005-0124885	A1	Abend et al.	06-09-2005
✓		2006-0205753	A1	Israel	09-14-2006

FOREIGN PATENT DOCUMENTS

Examiner's Initials	Cite No.	Foreign Patent Document			Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY	Translation (Y/N)
		Office/Country	Number	Kind Code			
PS		DE	196 51 551	A1	Klinge Pharm GmbH	06-18-1998	Y-Abstract
		WO	96/14058	A1	Euroceltique, S.A.	05-17-1996	
✓		WO	2004/091623	A1	Progenics Pharmaceuticals, Inc.	10-28-2004	

EXAMINER:

Phyllis Spivack

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9/12/07

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OTHER ART – NON PATENT LITERATURE DOCUMENTS

Examiner's Initials [#]	Cite No	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	Translation (Y/N)
PS		[No Author Listed] Ion Channels and Genetic Diseases, Chapter 1.	
		[No Author Listed] Remington's Pharmaceutical Sciences. 15 th Edition. 1995:1614-5.	
		[No Author Listed] Progenics initiates second phase 3 clinical trial of methylnaltrexone in opioid-induced constipation. Press Release. Progenics Pharmaceuticals, Inc. January 13, 2004.	
		[No Author Listed] Progenics achieves enrollment target in pivotal phase 3 clinical trial of methylnaltrexone for opioid-induced constipation. Press Release. Progenics Pharmaceuticals, Inc. December 3, 2004.	
		[No Author Listed] Progenics announces positive top-line results from pivotal phase 3 clinical trial of MNTX in opioid-induced constipation. Press Release. Progenics Pharmaceuticals, Inc. March 10, 2005.	
		AUNG et al., Methylnaltrexone prevents morphine-induced kaolin intake in the rat. Life Sci. 2004 Apr 16;74(22):2685-91.	
		BROWN et al., Techniques for mechanical stimulation of cells in vitro: a review. J Biomech. 2000 Jan;33(1):3-14.	
		EGAN et al., Prospective pharmacokinetic and pharmacodynamic validation of propofol's context sensitive T1/2. Anesthesiology. 1999 Sep;91(3A): Abstract A347.	
		FOSS et al., The efficacy of oral methylnaltrexone in decreasing the subjective effects of IV morphine. Anesth Analg. 1997;84. Abstract S484.	
		FOSS et al., Enteric-coated methylnaltrexone prevents opioid-induced oral-cecal transit delay in humans. Anesth Analg. 2000;90. Abstract S409.	
		FOSS et al., Subcutaneous methylnaltrexone reduces morphine-induced subjective effects in humans. Anesthesiology. 2001;95. Abstract A-817.	
		FRANCE et al., Morphine, saline and naltrexone discrimination in morphine-treated pigeons. J Pharm and Exper Ther. 1987;242:195-202.	
		FUNKE et al., A proton and carbon-13 nuclear magnetic resonance study of three quaternary salts of naloxone and oxymorphone. J Chem Soc. 1986:735-8.	
		GUTSTEIN et al., Role of inositol 1,4,5-trisphosphate receptors in regulating apoptotic signaling and heart failure. Heart Vessels. 1997;Suppl 12:53-7.	
		HICKS et al., Differential effects of the novel non-peptidic opioid 4-tyrosylamido-6-benzyl-1,2,3,4 tetrahydroquinoline (CGPM-9) on in vitro rat lymphocyte and macrophage functions. Life Sci. 2001 May 4;68(24):2685-94.	
		HIROTA et al., Loss of a gp130 cardiac muscle cell survival pathway is a critical event in the onset of heart failure during biomechanical stress. Cell. 1999 Apr 16;97(2):189-98.	
		HO et al., Beta-endorphin: peripheral opioid activity of homologues from six species. Int J Pept Protein Res. 1987 Apr;29(4):521-4.	
		HO et al., Methylnaltrexone antagonizes opioid-mediated enhancement of HIV infection of human blood mononuclear phagocytes. J Pharmacol Exp Ther. 2003 Dec;307(3):1158-62.	
✓		HUSSAIN et al., Improvement of the oral bioavailability of naltrexone in dogs: a prodrug approach. J Pharm Sci. 1987 May;76(5):356-8.	

EXAMINER:	DATE CONSIDERED:
Phyllis Spivack	9/12/07

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FILING DATE:	April 8, 2004	CONFIRMATION NO.:	9059
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DS	HUSSAIN et al., Naltrexone-3-salicylate (a prodrug of naltrexone): synthesis and pharmacokinetics in dogs. <i>Pharm Res.</i> 1988 Feb;5(2):113-5.	
11	IORIO et al., Narcotic agonist/antagonist properties of quaternary diastereoisomers derived from oxymorphone and naloxone. <i>Eur J Med Chem.</i> 1984;19(4):301-3.	
	LOPEZ et al., Demonstration of long-lasting blockade of experimental ileus in rats by an opioid kappa-agonist. <i>Gastroenterology.</i> 1995;108(4):A640. Abstract.	
	MCCARTHY et al., Preliminary studies on the use of plasma beta-endorphin in horses as an indicator of stress and pain. <i>J Equine Vet Sci.</i> 1993;13(4):216-9.	
	MIEDEMA et al., Methods for decreasing postoperative gut dysmotility. <i>Lancet Oncol.</i> 2003 Jun;4(6):365-72.	
	MOSS et al., Methylnaltrexone prevents morphine-induced CCR5 receptor expression. <i>Anesthesiology.</i> 2003;99. Abstract A-961.	
	PAPAPETROPOULOS et al., Nitric oxide synthase inhibitors attenuate transforming-growth-factor-beta 1-stimulated capillary organization in vitro. <i>Am J Pathol.</i> 1997 May;150(5):1835-44.	
	RESNICK et al., Delayed gastric emptying and postoperative ileus after nongastric abdominal surgery: part I. <i>Am J Gastroenterol.</i> 1997 May;92(5):751-62.	
	RESNICK et al., Delayed gastric emptying and postoperative ileus after nongastric abdominal surgery: part II. <i>Am J Gastroenterol.</i> 1997 Jun;92(6):934-40.	
	STEPHENSON et al., Methylnaltrexone reverses opioid-induced constipation. <i>Lancet Oncol.</i> 2002 Apr;3(4):202.	
	THOMAS et al., Amelioration of peripheral side effects of opioids: clinical experience with methylnaltrexone (MNTX). <i>Proc World Congr Anesth.</i> 2004;107.	
	TOMIYASU et al., Analysis of intercostal nerve damage associated with chronic post-thoracotomy pain. <i>Anesthesiology.</i> 2001;95. Abstract A-964.	
	WEI et al., Opioid-induced immunosuppression: is it centrally mediated or peripherally mediated? <i>Biochem Pharmacol.</i> 2003 Jun 1;65(11):1761-6.	
	WEI et al., Pharmacokinetics of subcutaneous methylnaltrexone: different route administration comparison. 2001. ASA Annual Meeting Abstracts. October 14-18, 2001. Chicago, IL. Abstract A-962.	
	YUAN et al., Gut and brain effects of American ginseng root on brainstem neuronal activities in rats. <i>Amer J Chin Med.</i> 1998; 26: 47-55.	
	YUAN et al., Methylnaltrexone (MNTX) for chronic opioid-induced constipation. 2002 ASCO Annual Meeting. <i>Proc Am Soc Clin Oncol.</i> 2002;21:376a. Abstract 1501.	
	YUAN et al., Safety and tolerance of oral methylnaltrexone in healthy volunteers. <i>Anesth Analg.</i> 1997;84:S1-599. Abstract S574.	
	YUAN et al., Methylnaltrexone changes gut motility and transit time in chronic methadone-maintained subjects. <i>Anesth Analg.</i> 1999;88: S1-424. Abstract S404.	
	YUAN et al., Antagonism of chronic opioid-induced gut effects. <i>Anesth Analg.</i> 2000;90:S1-523. Abstract S479.	
	YUAN et al., Pharmacokinetics of intravenous vs. oral methylnaltrexone: evidence for direct gut effects. <i>Anesth Analg.</i> 2001;92: S1-363. Abstract S274.	
↓	YUAN et al., Oral methylnaltrexone reverses morphine-induced changes in gastrointestinal motility. <i>Anesthesiology.</i> 1995 Sep;85(3A). Abstract A335.	

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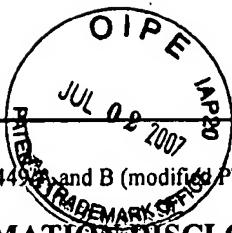
PS	YUAN et al., Oral methylnaltrexone reverses chronic opioid-induced constipation. Anesthesiology. 2000 Sep;93(3A). Abstract A-872.	
	YUAN et al., Subcutaneous methylnaltrexone prevents morphine-induced delay in gut transit time: a clinical trial. Anesthesiology. 2001;95. Abstract A-963.	
	YUAN et al., Methylnaltrexone prevents morphine-induced kaolin intake in the rat. Anesthesiology. 2003;99. Abstract A-922.	
	YUAN et al., Dose-related effects of oral acetaminophen on cold-induced pain: a double-blind, randomized, placebo-controlled trial. Clin Pharmacol Ther. 1998 Mar;63(3):379-83.	
	YUAN et al., Gastric effects of mu-, delta- and kappa-opioid receptor agonists on brainstem unitary responses in the neonatal rat. Eur J Pharmacol. 1996 Oct 24;314(1-2):27-32.	
	YUAN et al., Effects of low-dose morphine on gastric emptying in healthy volunteers. J Clin Pharmacol. 1998 Nov;38(11):1017-20.	
	YUAN et al., Gut motility and transit changes in patients receiving long-term methadone maintenance. J Clin Pharmacol. 1998 Oct;38(10):931-5.	
	YUAN et al., Tolerability, gut effects, and pharmacokinetics of methylnaltrexone following repeated intravenous administration in humans. J Clin Pharmacol. 2005 May;45(5):538-46.	
	YUAN et al., Antagonism of gastrointestinal opioid effects. Reg Anesth Pain Med. 2000 Nov-Dec;25(6):639-42.	
	YUAN et al., Methylnaltrexone reduces oral-cecal transit time in humans. Dig Dis Week Abstr. 2003:A-578. Abstract T1840.	
	YUAN et al., Opioid analgesia without gut side effects: effects of methylnaltrexone as a novel peripheral opioid antagonist. Assoc Univ Anesth Abst. 2003: PD2.	
✓	YUAN et al., Pain control with side effects: clinical studies on methylnaltrexone as a novel peripheral opioid antagonist. 7 th America-Japan Anesth Congr. Yamanashi, Japan. 2002:41.	

*a copy of this reference is not provided as it was previously cited by or submitted to the office in a prior application, Serial No. ___, filed ___, and relied upon for an earlier filing date under 35 U.S.C. 120 (continuation, continuation-in-part, and divisional applications).

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OTHER ART — NON PATENT LITERATURE DOCUMENTS

Examiner's Initials #	Cite No.	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	Translation (Y/N)
PS		ZIMMERMAN et al., Discovery of a potent, peripherally selective trans-3,4-dimethyl-4-(3-hydroxyphenyl)piperidine opioid antagonist for the treatment of gastrointestinal motility disorders. J Med Chem. 1994 Jul 22;37(15):2262-5.	

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